



PATENT

Docket No.: 20747/270 (ARDBW/P29257US)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant : Axel Thomson

Serial No. : 10/528,267

Cnfrm. No. : To Be Assigned

Filed : September 17, 2003

For : A METHOD OF PROTECTING A PATIENT  
FROM POSSIBLE ADVERSE EFFECTS OF  
INHIBITION OF THE SHH-SIGNALLING  
PATHWAY BY SUPPRESSING  
TESTOSTERONE OR ITS EFFECTS IN THE  
PATIENT

Examiner:  
To Be Assigned

Art Unit:  
To Be Assigned

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SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT  
UNDER 37 CFR §§ 1.97-1.98

Mail Stop: Amendment  
Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

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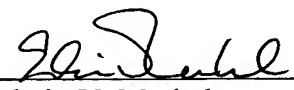
Pursuant to 37 CFR §§ 1.97-1.98, applicant hereby brings to the attention of the United States Patent and Trademark Office, the enclosed references listed on the attached PTO/SB/08 form.

Pursuant to 37 C.F.R. § 1.98(a)(2)(ii), copies of the cited U.S. Patents (i.e., Reference Cite Nos. 1-2) are not enclosed. Copies of all other listed references (i.e., Reference Cite Nos. 3-67) are enclosed herewith.

Pursuant to 37 CFR § 1.97(b)(3), no fee is required. If additional fees are required, however, the Commissioner is hereby authorized to charge any fees to Deposit Account No. 14-1138.

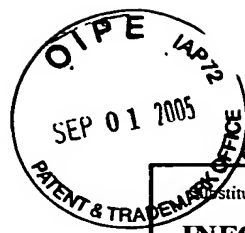
Respectfully submitted,

Date: August 29, 2005

  
Edwin V. Merkel  
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<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)				<b>Complete if Known</b>	
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				Filing Date	September 17, 2003
				First Named Inventor	Axel Thomson
				Art Unit	To Be Assigned
Examiner Name	To Be Assigned				
Sheet	1	of	6	Attorney Docket Number	20747/270 (ARDBW/P29257US)

U.S. PATENT DOCUMENTS					
Examiner Initials <sup>1</sup>	Cite No. <sup>1</sup>	U.S. Patent Document	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
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	1	US-4,946,779	08-07-1990	Kameda et al.	
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FOREIGN PATENT DOCUMENTS						
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		Country Code <sup>3</sup> Number <sup>4</sup> Kind Code <sup>5</sup> (if known)				
	3	WO 99/52534 A1	10-21-1999	Beachy et al.		
	4	WO 00/41545 A1	07-20-2000	Dudek		
	5	WO 01/27135 A2	04-19-2001	Beachy		
	6	WO 02/30462 A2	04-18-2002	Dudek et al.		
	7	WO 02/36818 A2	05-10-2002	Martin et al.		

OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS			
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Substitute for form 1449B/PTO <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> <i>(use as many sheets as necessary)</i>				<b>Complete if Known</b>	
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	11	BHATIA-GAUR et al., "Roles for <i>Nkx3.1</i> in Prostate Development and Cancer," <i>Genes Dev.</i> 13:966-977 (1999)		
	12	BITGOOD & McMAHON, " <i>Hedgehog</i> and <i>Bmp</i> Genes Are Coexpressed at Many Diverse Sites of Cell-Cell Interaction in the Mouse Embryo," <i>Dev. Biol.</i> 172:126-138 (1995)		
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	14	CHEN et al., "Inhibition of Hedgehog Signaling by Direct Binding of Cyclopamine to Smoothed," <i>Genes Dev.</i> 16:2743-2748 (2002)		
	15	CHEN et al., "Small Molecule Modulation of Smoothed Activity," <i>Proc. Natl. Acad. Sci. USA</i> 99(22):14071-14076 (2002)		
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	18	CHOMCZYNSKI et al., "Single-Step Method of RNA Isolation by Acid Guanidinium Thiocyanate-Phenol-Chloroform Extraction," <i>Anal. Biochem.</i> 162:156-159 (1987)		
	19	COLE et al., "The EBV-Hybridoma Technique and Its Application to Human Lung Cancer," <i>Monoclonal Antibodies and Cancer Therapy</i> , Alan R. Liss, Inc., pp.77-96 (1985)		
	20	COTE et al., "Generation of Human Monoclonal Antibodies Reactive with Cellular Antigens," <i>Proc. Natl. Acad. Sci. USA</i> 80:2026-2030 (1983)		
	21	CUNHA & CHUNG, "Stromal-Epithelial Interactions-I. Induction of Prostatic Phenotype in Urothelium of Testicular Feminized (Tfm/y) Mice," <i>J. Steroid Biochem.</i> 14:1317-1321 (1981)		
	22	CUNHA et al., "Smooth Muscle-Epithelial Interactions in Normal and Neoplastic Prostatic Development," <i>Acta Anat.</i> 155:63-72 (1996)		

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	23	EPSTEIN, J.I., "Nonspecific Granulomatous Prostatitis vs. Infectious Granulomas," in <i>Differential Diagnosis in Pathology: Urologic Disorders</i> , Section 3: Prostate, New York, Igaku-Shoin Ltd., pp.74-136 (1992)	
	24	FREEMAN et al., "A Comparison of Basal Cell Markers Used in the Prostate," <i>Histopathology</i> 40:483-494 (2002)	
	25	FROHMAN et al., "Isolation of the Mouse <i>Hox-2.9</i> Gene; Analysis of Embryonic Expression Suggests that Positional Information Along the Anterior-Posterior Axis is Specified by Mesoderm," <i>Development</i> 110:589-607 (1990)	
	26	GAO et al., "Conversion from a Paracrine to an Autocrine Mechanism of Androgen-Stimulated Growth during Malignant Transformation of Prostatic Epithelial Cells," <i>Cancer Res.</i> 61:5038-5044 (2001)	
	27	GENBANK ACCESSION NO. U43148 (1996)	
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	29	GENBANK ACCESSION NO. U84401 (1997)	
	30	GENBANK ACCESSION NO. AF316573 (2002)	
	31	GENBANK ACCESSION NO. AB007298 (2002)	
	32	GENBANK ACCESSION NO. M57609 (1994)	
	33	HARAGUCHI et al., "Unique Functions of Sonic Hedgehog Signaling During External Genitalia Development," <i>Development</i> 128:4241-4250 (2001)	
	34	HAYWARD et al., "Epithelial Development in the Rat Ventral Prostate, Anterior Prostate and Seminal Vesicle," <i>Acta Anat.</i> 155:81-93 (1996)	

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	35	HAYWARD et al., "Interactions Between Adult Human Prostatic Epithelium and Rat Urogenital Sinus Mesenchyme in a Tissue Recombination Model," <i>Differentiation</i> 63:131-140 (1998)	
	36	INCARDONA et al., "The Teratogenic <i>Veratrum</i> Alkaloid Cyclopamine Inhibits Sonic Hedgehog Signal Transduction," <i>Development</i> 125:3553-3562 (1998)	
	37	KIM et al., " <i>Nkx3.1</i> Mutant Mice Recapitulate Early Stages of Prostate Carcinogenesis," <i>Cancer Res.</i> 62:2999-3004 (2002)	
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	41	LAMM et al., "Mesenchymal Factor Bone Morphogenetic Protein 4 Restricts Ductal Budding and Branching Morphogenesis in the Developing Prostate," <i>Dev. Biol.</i> 232:301-314 (2001)	
	42	LAMM et al., "Sonic Hedgehog Activates Mesenchymal <i>Glil</i> Expression during Prostate Ductal Bud Formation," <i>Dev. Biol.</i> 249:349-366 (2002)	
	43	MORRISON et al., "Chimeric Human Antibody Molecules: Mouse Antigen-Binding Domains with Human Constant Region Domains," <i>Proc. Natl. Acad. Sci. USA</i> 81:6851-6855 (1984)	
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	45	NOGAWA et al., "Bud Formation Precedes the Appearance of Differential Cell Proliferation During Branching Morphogenesis of Mouse Lung Epithelium In Vitro," <i>Dev. Dyn.</i> 213:228-235 (1998)	
	46	OLUMI et al., "Carcinoma-Associated Fibroblasts Direct Tumor Progression of Initiated Human Prostatic Epithelium," <i>Cancer Res.</i> 59:5002-5011 (1999)	

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	47	ORLANDI et al., "Cloning Immunoglobulin Variable Domains for Expression by the Polymerase Chain Reaction," <i>Proc. Natl. Acad. Sci. USA</i> 86:3833-3837 (1989)		
	48	PARSONS et al., "p63 Protein Expression is Rare I Prostate Adenocarcinoma: Implications for Cancer Diagnosis and Carcinogenesis," <i>Urology</i> 58:619-624 (2001)		
	49	PATHI et al., "Comparative Biological Responses to Human Sonic, Indian, and Desert Hedgehog," <i>Mech. Dev.</i> 106:107-117 (2001)		
	50	PEPICELLI et al., "Sonic Hedgehog Regulates Branching Morphogenesis in the Mammalian Lung," <i>Curr. Biol.</i> 8:1083-1086 (1998)		
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	53	ROELINK et al., "Floor Plate and Motor Neuron Induction by <i>vhh-1</i> , a Vertebrate Homolog of <i>hedgehog</i> Expressed by the Notochord," <i>Cell</i> 76:761-775 (1994)		
	54	RUIZ I ALTABA et al., "Gli and Hedgehog in Cancer: Tumours, Embryos and Stem Cells," <i>Nat. Rev. Cancer</i> 2:361-372 (2002)		
	55	SCHNEIDER et al., "Targeted Disruption of the Nkx3.1 Gene in Mice Results in Morphogenetic Defects of Minor Salivary Glands: Parallels to Glandular Duct Morphogenesis in Prostate," <i>Mech. Dev.</i> 95:163-174 (2000)		
	56	STONE et al., "The Tumour-Suppressor Gene <i>Patched</i> Encodes a Candidate Receptor for Sonic Hedgehog," <i>Nature</i> 384:129-134 (1996)		
	57	SUGIMURA et al., "Keratinocyte Growth Factor (KGF) Can Replace Testosterone in the Ductal Branching Morphogenesis of the Rat Ventral Prostate," <i>Int. J. Dev. Biol.</i> 40:941-951 (1996)		
	58	SUGIMURA et al., "Morphogenesis of Ductal Networks in the Mouse Prostate," <i>Biol. Reprod.</i> 34:961-971 (1986)		

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	59	SUKEGAWA et al., "The Concentric Structure of the Developing Gut is Regulated by Sonic Hedgehog Derived from Endodermal Epithelium," <i>Development</i> 127:1971-1980 (2000)	
	60	TAKEDA et al., "Construction of Chimaeric Processed Immunoglobulin Genes Containing Mouse Variable and Human Constant Region Sequences," <i>Nature</i> 314:452-454 (1985)	
	61	THOMSON & CUNHA, "Prostatic Growth and Development are Regulated by FGF10," <i>Development</i> 126:3693-3701 (1999)	
	62	THOMSON et al., "Analysis of Growth Factor and Receptor mRNA Levels During Development of the Rat Seminal Vesicle and Prostate," <i>Development</i> 124:2431-2439 (1997)	
	63	THOMSON et al., "The Role of Smooth Muscle in Regulating Prostatic Induction," <i>Development</i> 129:1905-1912 (2002)	
	64	TOTTEN et al., "Microscopic Differential Diagnosis of Latent Carcinoma of Prostate," <i>Arch. Pathol.</i> 55:131-141 (1953)	
	65	WANG et al., "Cell Differentiation Lineage in the Prostate," <i>Differentiation</i> 68:270-279 (2001)	
	66	WINTER & MILSTEIN, "Man-made Antibodies," <i>Nature</i> 349:293-299 (1991)	
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Substitute for form 1449B/PTO				<b>Complete if Known</b>	
<b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b> (use as many sheets as necessary)				Application Number	10/528,267
				Filing Date	September 17, 2003
				First Named Inventor	Axel Thomson
				Group Art Unit	To Be Assigned
				Examiner Name	To Be Assigned
Sheet	2	of	6	Attorney Docket Number	20747/270 (ARDBW/P29257US)

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	11	BHATIA-GAUR et al., "Roles for <i>Nkx3.1</i> in Prostate Development and Cancer," <i>Genes Dev.</i> 13:966-977 (1999)		
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	23	EPSTEIN, J.I., "Nonspecific Granulomatous Prostatitis vs. Infectious Granulomas," in <i>Differential Diagnosis in Pathology: Urologic Disorders</i> , Section 3: Prostate, New York, Igaku-Shoin Ltd., pp.74-136 (1992)	
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	35	HAYWARD et al., "Interactions Between Adult Human Prostatic Epithelium and Rat Urogenital Sinus Mesenchyme in a Tissue Recombination Model," <i>Differentiation</i> 63:131-140 (1998)	
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	59	SUKEGAWA et al., "The Concentric Structure of the Developing Gut is Regulated by Sonic Hedgehog Derived from Endodermal Epithelium," <i>Development</i> 127:1971-1980 (2000)	
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